

# STIC-Biotech/ChemLib

**From:** Fredman, Jeffrey  
**Sent:** Monday, January 07, 2002 3:25 PM  
**To:** STIC-Biotech/ChemLib  
**Cc:** Lacourciere, Karen  
**Subject:** FW: Sequence search approval 09/975,123

I Approve.

Jeff Fredman

-----Original Message-----

**From:** Lacourciere, Karen  
**Sent:** Monday, January 07, 2002 11:24 AM  
**To:** Fredman, Jeffrey  
**Subject:** Sequence search approval 09/975,123

Jeff-

Could you approve this sequence search? Each sequence is an antisense targeted to the same gene. Thank-you!  
Karen

Please perform a sequence search in the commercial databases on the following sequences for 09/975,123. Please perform the search as a **length limited search, please limit the length of oligonucleotides to less than 100 nucleotides long:**

13-19, 21, 23-36, and 38-43

Thank-you!

*Karen A. Lacourciere Ph.D.*  
CM1 11D09 GAU 1635  
(703) 308-7523

**mailbox 11E12**

Searcher: \_\_\_\_\_  
Phone: \_\_\_\_\_  
Location: \_\_\_\_\_

Office: \_\_\_\_\_  
Clerical: \_\_\_\_\_  
Online time: \_\_\_\_\_

TYPE OF SEARCH:  
NA Sequences: 28  
AA Sequences: \_\_\_\_\_  
Structures: \_\_\_\_\_

Patent Family: \_\_\_\_\_  
Other: \_\_\_\_\_

VENDOR/COST (where applic.)

STN: \_\_\_\_\_

DIALOG: \_\_\_\_\_

Questel/Orbit: \_\_\_\_\_

DPLink: \_\_\_\_\_

WWW/Internet: \_\_\_\_\_  
Other (specify): \_\_\_\_\_

11/12/03 10:00 AM

Ref# 100921

3/12/03

11/12

## SEARCH REQUEST FORM

Scientific and Technical Information Center

First name \_\_\_\_\_ TANE ZARA Last name \_\_\_\_\_ 77572 Date \_\_\_\_\_ 8/12/03  
 Address \_\_\_\_\_ 1635 Filing Number \_\_\_\_\_ 6-5820 Serial Number \_\_\_\_\_ 09/975123  
 City \_\_\_\_\_ Denville Return Address \_\_\_\_\_ 11003 Form of Report Preferred \_\_\_\_\_ CD/DISK EMAIL  
 Tel # \_\_\_\_\_ 1181

If more than one search is submitted, please prioritize searches in order of need. \_\_\_\_\_

Please enter the subject matter of the search topic and describe in as much detail as possible the subject matter to be searched. Please use the search phrase to include key words, synonyms, acronyms, and part numbers and combine with the concept or name of the invention. Define any terms that may have a special meaning. Give examples of relevant citations, authors, etc. if known. Please attach a copy of the cover sheet, pertinent claims, and abstract.

Title of Invention \_\_\_\_\_ Freezer

Inventor's primary occupation \_\_\_\_\_ AS & ILGFBP5

Earliest Priority Filing Date \_\_\_\_\_ 10/9/01

\*For Sequence Searches Only \* Please include all pertinent information (parent, child, divisional, or issued patent numbers) along with the appropriate serial number.

Please Search Seg ID #'s:  
 ✓ 15, ✓ 23, ✓ 24, 26 - 30  
 33 - 36, 38 - 43

✓  
 1181/03

For Interference & regular data bases.

— Limit ALL to 100 NT's. ONLY.

March.

\* These seg have been searched together before.

## STAFF USE ONLY

SLP/par

## Type of Search

## Vendors and cost where applicable

NA Sequence #

NA Interference

Structural #

Biological #

9/12/03

S2	1 Items	Description
S1	12	IGFBP1 OR IGF-1 W/ RIB
S2	13	SI AND ANTISENSE OR RIBONUCL
S3	13	R2 (unique items)
S4	14	SI AND ANTISENSE OR RIBONUCL
S5	14	R2 (unique items)
S6	14	INSULIN (W) LIKE (W) GROWTH (W) FACTOR (W) BINDING (W) PRO- TEIN (W) 5
S7	11	S6 (S) (ANTISENSE OR RIBONUCL)
S8	14	R2 (unique items)
S9	14	S6 AND (ANTISENSE OR RIBONUCL)
S10	14	R2 (unique items)

>>>KWIC option is not available in file S: 41, "S", See

10/3,K/1 (Item 1 from file: 5)

DIALOG(R)File 5: Biosis Previews(R)

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13216983 BIOSIS NO.: 200100424132

**The IGF/IGFBP system in CNS malignancy.**

AUTHOR: Zumkeller W(a); Westphal M

AUTHOR ADDRESS: (a)Department of Pediatrics, Martin-Luther-University  
Halle-Wittenberg, University Hospital, Ernst-Grube-Str. 40, 06097,  
Halle/Saale: walter.zumkeller@medizin.uni-halle.de\*\*Germany

JOURNAL: Molecular Pathology 54 (4):p227-229 August, 2001

MEDIUM: print

ISSN: 1366-8714

DOCUMENT TYPE: Article

RECORD TYPE: Abstract

LANGUAGE: English

SUMMARY LANGUAGE: English

...ABSTRACT: Both types of IGF receptor are expressed in gliomas and, in particular, the type I IGF receptor appears to be upregulated in malignant brain tissue. \*Antisense\* IGF-I receptor mRNA induces an antitumour response, resulting in complete brain tumour regression. Clinical trials for the treatment of brain tumours in humans based on a gene transfer protocol using IGF-I receptor \*antisense\* are under way. All six IGFBPs are expressed to a variable extent in brain tumours. High concentrations of IGFBP-2 are found in cerebrospinal fluid...

DESCRIPTORS:

CHEMICALS & BIOCHEMICALS: \*antisense\* insulin-like growth factor-I receptor messenger RNA...

...\*insulin\*--\*like\* \*growth\* \*factor\* \*binding\* \*protein\*--\*5\*;

10/3,K/2 (Item 2 from file: 5)

DIALOG(R)File 5: Biosis Previews(R)

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13216983 BIOSIS NO.: 200100424133

Castration-induced up-regulation of \*insulin\*--\*like\* \*growth\* \*factor\* \*binding\* \*protein\*--\*5\* potentiates insulin-like growth factor-I activity and accelerates progression to androgen independence in prostate cancer models.

AUTHOR: Miyake Hideaki; Pollak Michael; Gleave Martin E(a)

AUTHOR ADDRESS: (a)Division of Urology, University of British Columbia,  
19-2, 2254 Health Sciences, Vancouver, BC, V6Z 2B5\*\*Canada

JOURNAL: Cancer Research 61 (15):6171-6176 July 1, 2001

MEDIUM: print

Castration-induced up-regulation of \*insulin\*--\*like\* \*growth\* \*factor\*

\*binding\* \*protein\*\*5\* potentiates insulin-like growth factor-I activity and accelerates progression to androgen independence in prostate cancer models.

ABSTRACT: Although insulin-like growth factor binding protein-5 (IGFBP-5) has been shown to be implicated in prostate cancer progression, the functional role of IGFBP-5 in progression to androgen-independence remains largely...

...cells were stably transfected with IGFBP-5 gene, and IGFBP-5-over-expressing LNCaP tumors progressed significantly faster to androgen independence after castration compared with controls. Antisense nucleic acid oligoribonucleotides (ODNs) were then designed that reduced IGFBP-5 expression in Shionogi tumor cells in vitro in a dose-dependent and sequence-specific manner. Growth of Shionogi tumor cells was inhibited by antisense IGFBP-5 ODN treatment in a time- and dose-dependent manner, which could be reversed by exogenous IGF-I. However, antisense IGFBP-5 ODN treatment had no additive inhibitory effect on Shionogi tumor cell growth when IGF-I activity was neutralized by anti-IGF-I antibody. Antisense IGFBP-5 ODN treatment resulted in decreased mitogen-activated protein kinase activity and number of cells in the S + G2-M phases of the cell cycle that directly correlated with reduced proliferation rate of Shionogi tumor cells. Systemic administration of antisense IGFBP-5 ODN in mice bearing Shionogi tumors after castration significantly delayed time to progression to androgen independence and inhibited growth of AI recurrent tumors...  
...serves to enhance IGF bioactivity and raise the possibility that the response of prostate cancer to androgen withdrawal can be enhanced by strategies, such as antisense IGFBP-5 ODN therapy, that target IGF signal transduction.

DESCRIPTIONS:

CHEMICALS & BIOCHEMICALS: ...\*insulin\*\*like\* \*growth\* \*factor\* \*binding\* \*protein\*\*5\*--...

...human IGFBP-5 gene (human \*insulin\*\*like\* \*growth\* \*factor\* \*binding\* \*protein\*\*5\* gene) (Hominidae)

10/3,K/3 (Item 3 from file: 5)

DIALOG(R)File 5:Biosis Previews(R)  
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12099750 BIOSIS NO.: 199900394606

Inhibition of insulin-like growth factor I receptor signaling by the vitamin D analogue EB1089 in MCF-7 breast cancer cells: A role for insulin-like growth factor binding proteins.

AUTHOR: Rosen Florence; Pollak Michael(a)

AUTHOR ADDRESS: (a)Lady Davis Institute for Medical Research of the Jewish General Hospital, 3755 Cote Ste. Catherine, Canada

JOURNAL: International Journal of Oncology 15 (3):p589-594 Sept., 1999

ISSN: 0253-372X

ISI SOURCE: SCOPUS

RETRIEVE BY: ISI, SCOPUS

KEYWORD: EB1089

SUMMARY LANGUAGE: English

...ABSTRACT: : IBS-1 induced by insulin, IGF-I, and IGF-I analogue with greatly reduced affinity for IGFBI's. Furthermore, we demonstrate that an antisense IGFBP-5 oligoribonucleotide attenuates EB1089-induced inhibition of IGF-I-stimulated tyrosine phosphorylation of IBS-1 and ERK1/2. IGFBI's are required. This is, interestingly...

RETRIEVE BY:

12099750 BIOSIS NO.: 199900394606

12099750 BIOSIS NO.: 199900394606

12099750 BIOSIS NO.: 199900394606

1142292 BIOSIS NO.: 199800209631

Differential expression and localization of IGF-I and IGF binding proteins in inflamed rat colon.

AUTHOR: Jeek Isberg M; Mahapatra Niru; Land F; Kay; Eysselein Viktor E; McRoberts James A

AUTHOR ADDRESS: Harbor-UCLA Med. Cent., Div. Gastroenterol., Torrance, CA

••USA

JOURNAL: Journal of Receptor and Signal Transduction Research 18 (4-6):p 265-280 July-Nov., 1998

ISSN: 1079-9693

DOCUMENT TYPE: Article

RECORD TYPE: Abstract

LANGUAGE: English

...ABSTRACT: were sacrificed at 7 days after induction of colitis. Cryostat sections of colon from TNB-treated and control rats were hybridized with <sup>35</sup>S-labeled \*antisense\* probes for IGF-I $\kappa$ , IGFBP-3, IGFBP-4 and IGFBP-5. IGF-I mRNA was up-regulated in lamina propria cells, submucosa and smooth muscle...

DESCRIPTORS:

CHIMICALS & BIOCHEMICALS: ...\*insulin\*-\*like\* \*growth\* \*factor\* \*binding\* \*protein\*-\*5\*--

10/3,K/5 (Item 5 from file: 5)

DIALOG(R)File 5:Biosis Previews(R)  
(c) 2002 BIOSIS. All rts. reserv.

1142292 BIOSIS NO.: 199800209624

Up-regulation of \*insulin\*-\*like\* \*growth\* \*factor\* \*binding\* \*protein\*-\*5\* is independent of muscle cell differentiation, sensitive to rapamycin, but insensitive to wortmannin and LY294002.

AUTHOR: Rousse Sophie; Montarras Didier; Finset Christian; Dukcis Catherine

(a)

AUTHOR ADDRESS: (a) Inst. Natl. Sante Recherche Med., U.142, Hop. Saint Antoine, 75571 Paris Cedex 12\*\*France

JOURNAL: Endocrinology 139 (4):p1487-1495 April, 1998

ISSN: 0013-7227

DOCUMENT TYPE: Article

RECORD TYPE: Abstract

LANGUAGE: English

Up-regulation of \*insulin\*-\*like\* \*growth\* \*factor\* \*binding\* \*protein\*-\*5\* is independent of muscle cell differentiation, sensitive to rapamycin, but insensitive to wortmannin and LY294002.

...ABSTRACT: and modulated by IGF binding proteins (IGFBPs) secreted by the cells. The mouse C2 myoblast cell line stably transfected with a vector expressing IGF-II \*antisense\* RNA was used to show that specific IGFBP expression changes with the state of the cells: high levels of IGFBP-1 positive RNA when...

DESCRIPTORS:

CHIMICALS & BIOCHEMICALS: ...\*insulin\*-\*like\* \*growth\* \*factor\* \*binding\* \*protein\*-\*5\*--

10/3,K/6 (Item 6 from file: 5)

DIALOG(R)File 6:Biosis Previews(R)  
(c) 2002 BIOSIS. All rts. reserv.

ICARAI: H. et al. *Postgrad Med J* 1997; 73(866): 47-51. Cited in 1997  
CONFERENCE MEETING: 6th Joint Meeting of the European Society for  
Paediatric Endocrinology and the Lawson Wilkins Society for Pediatric  
Endocrinology, in collaboration with the Australian Paediatric Endocrine  
Group, the Japanese Society for Pediatric Endocrinology and the Latin  
American Society for Pediatric Endocrinology. Stockholm, Sweden. June  
22-26, 1997

ISSN: 0031-3025

RECORD TYPE: Citation

LANGUAGE: English

MISCELLANEOUS TERMS: ...\*INSULIN-LIKE GROWTH FACTOR BINDING PROTEIN-5\*  
\*ANTISENSE\* OLIGODEOXYNUCLEOTIDE...  
...\*INSULIN-LIKE GROWTH FACTOR BINGING PROTEIN-4\* \*ANTISENSE\* OLIGONUCLÉOTIDE  
...  
...\*INSULIN\*-\*LIKE\* \*GROWTH\* \*FACTOR\* \*BINDING\* \*PROTEIN\*-\*5\*;

10/3,K/7 (Item 7 from file: 5)  
DIALOG(R)File 5:Biosis Previews,RT  
(c) 2002 BIOSIS. All rts. reserv.

10700630 BIOSIS NO.: 199799321775

A role for \*insulin\*-\*like\* \*growth\* \*factor\* \*binding\* \*protein\* \*5\* in  
the antiproliferative action of the antiestrogen ICI 182780.

AUTHOR: Hlynch Hung(a); Yang Xiao-Feng; Pollak Michael

AUTHOR ADDRESS: (a)Dep. Med., McGill Univ., 3755 Cote Ste Catherine Rd.,  
Montreal, PQ H3T 1E2\*\*Canada

JOURNAL: Cell Growth & Differentiation 7 (11):p1501-1506 1996

ISSN: 1044-9523

RECORD TYPE: Abstract

LANGUAGE: English

A role for \*insulin\*-\*like\* \*growth\* \*factor\* \*binding\* \*protein\* \*5\* in  
the antiproliferative action of the antiestrogen ICI 182780.

...ABSTRACT: mRNA abundance, and increased IGFBP-5 protein accumulation in  
the conditioned medium. Growth stimulation following estradiol exposure  
was associated with opposite effects. An IGFBP-5 \*antisense\*  
oligodeoxynucleotide significantly decreased IGFBP-5 accumulation in  
conditioned media and enhanced MCF-7 cell DNA synthesis. Furthermore,  
this \*antisense\* oligodeoxynucleotide attenuated both  
antiestrogen-induced IGFBP-5 accumulation and antiestrogen-induced growth  
inhibition. These data demonstrate that estradiol down-regulates and ICI  
up-regulates an...

MISCELLANEOUS TERMS: ...\*INSULIN\*-\*LIKE\* \*GROWTH\* \*FACTOR\* \*BINDING\*  
\*PROTEIN\* \*5\*; \*INSULIN\*-\*LIKE\* \*GROWTH\* \*FACTOR\* \*BINDING\* \*PROTEIN\*  
\*5\* \*mRNA\*

10/3,K/8 (Item 8 from file: 5)

DIALOG(R)File 5:Biosis Previews,RT  
(c) 2002 BIOSIS. All rts. reserv.

108-4130 BIOSIS NO.: 199799321776

Osteogenic protein-1-mediated insulin-like growth factor gene expression in  
primary cultures of rat osteoblastic cells.

AUTHOR: Yeh Lin-Chuan C. M.; Adam Martin L; Kitten Allison M; Olson Merle J  
; ...

AUTHOR ADDRESS: Dept. Biochem., Univ. Texas Health Sci. Ctr., San Antonio, TX 78229

...ABSTRACT: a 10-kDa protein in myoblasts. The IGF-I- $\alpha$ , - $\beta$ , and - $\gamma$  mRNA levels decreased dramatically in an IGF-concentration-dependent manner. In addition, synthesis of 'antisense' oligoribonucleotides corresponding to IGF-I  $\alpha$ -III mRNA sequence with 5'-I reduced the IGF-I-induced elevation in alkaline phosphatase activity. The present results...  
MISCELLANEOUS TERMS: ...\*INSULIN\*-\*IGF\* \*GROWTH\* \*FACTOR\*-\*BINDING\*  
\*PROTEIN\*-\*5\*;

10/3,K/9 (Item 9 from file: 5)

DIALOG(R)File 5:Biosis Previews(R)  
(c) 2002 BIOSIS. All rts. reserv.

10381965 BIOSIS NO.: 199699004110

\*Insulin\*-\*like\* \*growth\* \*factor\* \*binding\* \*protein\*-\*5\* modulates muscle differentiation through an insulin-like growth factor-dependent mechanism.

AUTHOR: James Payton L; Stewart Claire E H; Rotwein Peter

AUTHOR ADDRESS: Dep. Biochemistry Molecular Biophysics, 660 South Euclid Ave., Box 8261, St. Louis, MO 63110\*\*USA

JOURNAL: Journal of Cell Biology, 133 (3):p683-693 1996

ISSN: 0021-9525

DOCUMENT TYPE: Article

RECORD TYPE: Abstract

LANGUAGE: English

\*Insulin\*-\*like\* \*growth\* \*factor\* \*binding\* \*protein\*-\*5\* modulates muscle differentiation through an insulin-like growth factor-dependent mechanism.

...ABSTRACT: sense myoblasts show enhanced survival in low serum medium, remaining viable for at least four weeks in culture. By contrast, myoblasts expressing the IGFBP-5 \*antisense\* transcript differentiate prematurely and more extensively than control cells. The inhibition of myogenic differentiation by high level expression of IGFBP-5 could be overcome by...

10/3,K/10 (Item 10 from file: 5)

DIALOG(R)File 5:Biosis Previews(R)  
(c) 2002 BIOSIS. All rts. reserv.

09535075 BIOSIS NO.: 199598049993

Localization of messenger ribonucleic acid for insulin-like growth factor-binding proteins in human skin by in situ hybridization.

AUTHOR: Hatch J A(a); Merruri F A; Edmondson S R; Werther G A

AUTHOR ADDRESS: (a)Cent. Hormone Res., Royal Children's Hosp., Flemington Rd., Parkville, 3052 VIC\*\*Australia

JOURNAL: Journal of Clinical Endocrinology & Metabolism, 134 (5):p1444-1449 1994

ISSN: 0021-9724

DOCUMENT TYPE: Article

RECORD TYPE: Abstract

LANGUAGE: English

...ABSTRACT: In an attempt to study the distribution of IGF-BPs in human skin, we have used in situ hybridization to localize messenger ribonucleic acid (mRNA) for the six IGF-BPs. 'Antisense' and 'sense' RNA probes for the IGF-BPs IGFBP-1, -2, -3, -4, -5, and -6 were prepared, and  $^{35}$ S-UTP sections of normal adult human skin were studied...  
MISCELLANEOUS TERMS: ...\*IGFBP\*-\*IGF\* \*GROWTH\* \*FACTOR\*-\*BINDING\*

Increased expression of IGF-binding protein-5 in Duchenne muscular dystrophy (DMD) fibroblasts correlates with the fibroblast-induced downregulation of DMD myoblast growth: an in vitro analysis.

Melone M A; Bellinc S; Gaidetisli S; Bellinc S; Istritofa R  
Second Division of Neurology, Second University of Naples, School of  
Medicine, Naples, Italy. marilena.melone@unipa.it

Document type: Standard

JOURNAL OF CLIMATE

第三章 亂世之亂世：明末清初

卷之三十一

... (GFPP-1) in DMD fibroblast-conditioned media by means of specific antibodies, or inhibiting GFPP-1 gene expression in DMD fibroblasts by means of antisense, the fibroblast-conditioned media lost inhibitory power over DMD myoblast proliferation. Copyright 2003 Wiley-Liss, Inc.

Descriptors: Fibroblasts--metabolism--MF; \*Fibroblasts--pathology--PA; \*Insignin-Like\* \*Growth-Factor--Binding--\*Protein\* \*5--biosynthesis--BI; \*Muscle, Skeletal--metabolism--MF; \*Muscle, Skeletal--pathology--PA; \*Muscular Dystrophy, Duchenne--metabolism--MF; \*Muscular Dystrophy, Duchenne--pathology--PA

Chemical Name: Culture Media, Conditioned; \*Insulin\*-Like\* \*Growth\*-  
\*Factor\*-\*Binding\*-\*Protein\* \*5\*

10/3.K/12 (Item 2 from file: 155)

DIALOG(R) File 155; MEDLINE(R)

08764041 96109186 PMTD: 8618825

Insulin-like growth factor II mediates epidermal growth factor-induced mitogenesis in cervical cancer cells.

Stellier M A; Delgado C H; Zou Z  
Section of Gynecologic Oncology, National Cancer Institute, Bethesda, MD

20892-1302, USA.  
Proceedings of the National Academy of Sciences of the United States of America (UNITED STATES) Dec 19 1995, 92 (26) p11970-4, ISSN 0027-8424

Journal Code: 7505876

Document type: Jou

Languages: ENGLISH

Main Citation Owner: N

... induced mitogenesis was abrogated in a dose-dependent manner by IGF-binding protein 5 (IGFBP-5), which binds to IGF-II and neutralizes it. An "antisense" oligonucleotide to TGF- $\beta$ 1 also inhibited the proliferative response to EGF. In addition, prolonged, but not short-term, stimulation with EGF required the presence of small amounts of

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Expression of the genes encoding the insulin-like growth factors (IGF-I and II), the IGF and insulin receptors, and IGF-binding proteins-1-6 and the localization of their gene products in normal and polycystic ovary syndrome ovaries.

el-Kerisy A; Chen M; Roberts M J; Shimakawa S; Ling N; LeRoith D; Roberts C M; Yen PM

Department of Reproductive Medicine, University of California School of Medicine, Los Angeles 90095.

Journal of Clinical Endocrinology and Metabolism (UNITED STATES) Jan 1991, Vol 131, No 1, p147-56, ISSN 0021-9724, Journal Code: 0375362

Document Type: Journal Article

Languages: ENGLISH

## Main Station, Sweden: NM

Record type: Completed

... in specific cellular compartments of normal and PCOS human ovaries. Messenger ribonucleic acid (mRNA) was localized by *in situ* hybridization with specific 35S-labeled human \*antisense\* RNA probes, and protein was detected by immunohistochemistry using specific antisera. Thecal cells, but not granulosa cells (GC), of small antral follicles (3-6 mm...

...; Growth Factor II--analysis--AN; Insulin-Like Growth Factor-Binding Protein 2; Insulin-Like Growth-Factor-Binding Protein 1; Insulin-Like Growth-Factor-Binding Proteins; \*Insulin\*-Like\* \*Growth\*-Factor\*-Binding\*-Protein\* '3'; Insulin-Like-Growth Factor Binding Protein 6; Middle Age; Ovary--pathology--PA; Polycystic Ovary Syndrome--pathology--PA; RNA Probes; Receptor, IGF Type I--analysis--AN...

...Chemical Name: Growth-Factor-Binding Protein 4; Insulin-Like Growth Factor-Binding Protein 2; Insulin-Like Growth-Factor Binding Protein 1; Insulin-Like Growth-Factor-Binding Proteins; \*Insulin\*-Like\* \*Growth\*- \*Factor\*-Binding\*-\*Protein\* \*5\*; Insulin-Like-Growth-Factor-Binding Protein 6; RNA Probes; RNA, Messenger; Receptor, IGF Type 2; Insulin-Like Growth Factor I; Insulin-Like Growth Factor II...

10/3, K/14 (Item 1 from file: 399)

DIALOG(E) File 399:CA SEARCH(R)

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136321287 CA: 136(21)321287w PATENT

Use of pregnancy-associated plasma protein-A2 (PAPP-A2), a novel insulin-like growth factor-binding protein-5 proteinase, for diagnosis and treatment of fetal abnormalities

LOCATION: Ben-

ASSIGNEE: Geno Biotech, Inc.

Patent: 20090294255 Date: 2009-11-26

<sup>10</sup> See, for example, the discussion of the 1992 Constitutional Convention in the *Constitutional Convention of 1992: The Final Report* (1993).

Use of antisense nucleic acids/analogs inhibiting growth factor-mediated cell proliferation for treatment of proliferative and/or inflammatory skin disorders

INVENTOR/AUTHOR: Werner, George Arthur; Wright, Christopher John  
LOCATION: Australia

ASSIGNEE: Royal Children's Hospital Research Foundation

PATENT: PCT International; WO 92/06641 DATE: 06/11/1992

APPLICATION: WO 91/04441. 471CC (AU 840113 (1943115))

PAGES: 111 pp. JURIS: FINNLAND LANGUAGE: English CLASS: A61K-31/00; A61K-31/12; C07K-17/14B; C12N-1/01/14B DESIGNATED COUNTRIES: AM; AT; AU; BE; BG; BR; BY; CA; CH; CN; DK; DE; ES; FI; GB; IE; HU; IS; JP; KE; KG; KP; KR; KE; LK; LR; LT; LV; MN; MG; MN; MM; NO; NZ; PL; PT; RO; RU; SD; SE; SG; SI; SK; TR; TM; TT DESIGNATED REGIONAL: KE; MW; SE; SZ; IS; AT; BE; CH; DE; DK; ES; FR; GB; GR; IT; LV; MG; NL; PT; SE; BF; BJ; CF; CG; CI; CM; GA; GU; ML; MR; NE; SN; TD; TG

10/3,K/16 (Item 1 from file: 35)

DIALOG(R)File 159:Dissertation Abs online  
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01439558 ORDER NO: AADAA-19833739

\*INSULIN\*-\*LIKE\* \*GROWTH\* \*FACTOR\* \*BINDING\* \*PROTEIN\*-\*5\* INHIBITS  
MYOGENIC DIFFERENTIATION THROUGH AN IGF-DEPENDENT PROCESS

Author: JAMES, PAYTON LEIGH

Degree: PH.D.

Year: 1995

Corporate Source/Institution: WASHINGTON UNIVERSITY (0252)

Source: VOLUME 56/06-B OF DISSERTATION ABSTRACTS INTERNATIONAL.

PAGE 2987. 168 PAGES

\*INSULIN\*-\*LIKE\* \*GROWTH\* \*FACTOR\* \*BINDING\* \*PROTEIN\*-\*5\* INHIBITS  
MYOGENIC DIFFERENTIATION THROUGH AN IGF-DEPENDENT PROCESS

...dependent manner.

Stable transfectants of the C2 cell line were established which constitutively express the coding sequence of IGFBP-5 in either the sense or \*antisense\* orientation to determine the function of this protein during myogenic differentiation. Forced expression of the \*antisense\* transcripts caused rapid differentiation as assessed by myotube formation, creatine kinase activity, and the production of myosin heavy chain and the muscle-specific transcription factor...

10/3,K/17 (Item 1 from file: 159)

DIALOG(R)File 159:Cancerlit  
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02322558 PMID: 97604744

Antiproliferative effects of ICI 182780 are partly mediated by upregulation of \*insulin\*-\*like\* \*growth\* \*factor\* \*binding\* \*protein\* \*5\* (IGFBP-5) (Meeting abstract).

Bray, J.; Lin, J.; Li, J.; Li, J.

Cancer Res. Research Inst., Monash U., Victoria 3168, Australia  
Int. J. Cancer Res. Relat. Biol. Eff. 1997, 13, 1119-1124

Document Type: MEETING ABSTRACT

Language: ENGLISH

Main Citation Owner: NCI/NIN

Report type: Completed

Antiproliferative effects of ICI 182780 are partly mediated by upregulation of \*insulin\*-\*like\* \*growth\* \*factor\* \*binding\* \*protein\* \*5\*

and anti-insulin antibodies to insulin. These results indicate that the beta cell releases anti-insulin loops and insulin at IGFII-1...

Chemical Name: IGF-I (1-70); Secretin; Histidine Antagonists; Insulin-Like Growth Factor-1; Insulin-Activating Protein-1; RNA, Messenger

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5/3, K/1 (Item 1 from file: 5)

MONITORING: S. P. L. S. PRELIMINARY  
MONITORING, ALL R.S. REPORT.

1990-1991 1991-1992 1992-1993 1993-1994

## Mesenchymal-epithelial transition in the developing metanephric kidney:

## Gene expression study by differential display.

AUTHOR: Filsov Sergei I(a); Ivanov Sergey V; Yoshino Kiyoshi; Dove Lee F;  
Filsova Tatiana M; Sijinathan Kathleen S; Karavanova Irina; Lerman  
Michael; Purohit Alash S

**AUTHOR ADDRESS:** National Cancer Institute-FCRU, Bldg. 338, Room 205E,  
Frederick, MD, 21702-1201, USA

JOURNAL: *Genesis: The Journal of Genetics and Development* 27 (1):p22-31

May, 2000

METHYL  
METHACRYLATE

ISSN: 1526-954X

DOCUMENT TYPE: Article

RECORD TYPE: Abst

LANGUAGE: English

...ABSTRACT: proteins (SH3-domain binding protein, G-protein-coupled receptor, Ser-Thr protein kinase), cell adhesion molecules (syndecan-4, integrin-beta1), and also gene33, H19, SM20, \*IGFBP5\*, MAMA receptor, lectin, keratin, beta-tubulin, calreticulin, GRP70, ERP72, MnSOD, thioredoxin, and others. Some have previously been associated with kidney development and serve as good controls for expected changes, while most have not been linked with kidney epithelial cell differentiation. Using thin sections of embryonic kidney and labeled \*antisense\* RNA probes, we applied RNA hybridization to confirm the results of DD and related the expression of these genes to specific cell lineages of the...

5/3.K/2 (Item 2 from file: 5)

DIALOG(R)File 5:Biosis Previews(R)  
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11428792 BIOSIS NO : 199800209624

Up-regulation of insulin-like growth factor binding protein-5 is independent of muscle cell differentiation, sensitive to rapamycin, but insensitive to wortmannin and LY294002

In-sensitive to wortmannin and Li294002.

<sup>10</sup> See, for example, the discussion of the 1992 Constitutional Convention in the *Constitutional Convention of 1992: The Final Report* (1993).

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2020年1月1日-2020年1月31日

## REVIEW ARTICLE: *Wittgenstein and the Philosophy of Mathematics* by Michael Friedman

### REFERENCES: *Continued*

5/3,K/3 (Item 1 from file: 399)

DIALOG(R) File 399:CA SEARCH(E)

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136321287 CA: 136(21)321287w PATENT

Use of pregnancy-associated plasma protein-A2 (PAPP-A2), a novel insulin-like growth factor-binding protein-5 proteinase, for diagnosis and treatment of fetal abnormalities

INVENTOR(AUTHOR): (xvi); Claus; Overgaard, Michael Tett

LOCATION: Den.

ASSIGNEE: Ceme Biotech Aps

PATENT: PCT International ; WO 200201953 A2 DATE: 20020426

APPLICATION: WO 200105695 (20011019) \*PCT 20001831 (20001020) \*US PV241840

APPLICATION

PAGES: 113 pp. CODEN: PIXMD2 LANGUAGE: English CLASS: C07K-014/435A  
DESIGNATED COUNTRIES: AE; AG; AL; AM; AT; AU; AZ; BA; BB; BG; BR; BY; BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DK; DM; DZ; EC; EE; ES; FI; HU; ID; IL; IN; IS; JP; KE; KG; KP; KR; KZ; LC; LK; LR; LS; LT; LU; LV; MA; MD; MG; MK; MN; MW; MX; MZ; NO; NZ; PH; PL; PT; RO; RU; SD; SE; SG; SI; SK; SL; TJ; TM; TR; TT; TZ; UA; UG; US; UZ; VN; YU; ZA; ZW; AM; AZ; BY; KG; KZ; MD; RU DESIGNATED REGIONAL: GH; GM; KE; LS; MN; NZ; SD; SL; SZ; TZ; UG; ZW; AT; BE; CH; CY; DE; DK; ES; FI; FR; GB; GR; IE; IT; LU; MC; NL; PT; SE; TR; BF; BJ; CF; CI; CM; GA; GN; GQ; GW; XI; MR; NE; SN; TD; TG

5/3,K/4 (Item 2 from file: 399)

DIALOG(R) File 399:CA SEARCH(E)

(c) 2002 AMERICAN CHEMICAL SOCIETY. All rts. reserv.

134125934 CA: 134(10)125934z PATENT

IGFBP-5 antisense oligodeoxynucleotide therapy for hormone-regulated tumors

INVENTOR(AUTHOR): Gleave, Martin

LOCATION: Can.,

ASSIGNEE: The University of British Columbia; Miyake, Hideaki

PATENT: PCT International ; WO 200105435 A2 DATE: 20010125

APPLICATION: WO 2000CA853 (20000719) \*US PV144495 (19990719)

PAGES: 45 pp. CODEN: PIXXR2 LANGUAGE: English CLASS: A61K-041/0CA

DESIGNATED COUNTRIES: AE; AG; AL; AM; AT; AU; AZ; BA; BB; BG; BR; BY; BZ; CA; CH; CN; CR; CU; CZ; DE; DK; DM; DZ; EC; EE; FI; GB; GD; GE; GH; GM; HR; HU; ID; IL; IN; IS; JP; KE; KG; KP; KR; KZ; LC; LK; LR; LS; LT; LU; LV; MA; MD; MG; MK; MN; MW; MX; MZ; NO; NZ; PL; PT; RO; RU; SD; SE; SG; SI; SK; SL; TJ; TM; TR; TT; TZ; UA; UG; US; UZ; VN; YU; ZA; ZW; AM; AZ; BY; KG; KZ; MD; RU; TJ; TM DESIGNATED REGIONAL: GH; GM; KE; LS; MW; MZ; SD; SL; SZ; TZ; UG; ZW; AT; BE; CH; CY; DE; DK; ES; FI; FR; GB; GR; IE; IT; LU; MC; NL; PT; SE; TR; BF; BJ; CF; CI; CM; GA; GN; GQ; GW; XI; MR; NE; SN; TD; TG

3/3,K/1 (Item 1 from file: 5)  
DIALOG(R)File 5:Biosis Previews(R)  
(c) 2002 BIOSIS. All rts. reserv.  
11428292 BIOSIS NO.: 11428292  
Mesenchymal-epithelial transition in the developing metanephric kidney:  
Gene expression study by differential display.

AUTHOR: Plisova Tatjana V; Ivanov Svetozar V; Yashini Kiyoshi; Sung Lee H;  
Plisova Tatjana V; Hightower Kathleen S; Karavanova Irina; Lerman  
Michael; Perantoni Alan C  
AUTHOR ADDRESS: N.I. National Cancer Institute-MKRC, Bldg. 37C, Rm. 2C14,  
Frederick, MD, 21702-1217 USA  
JOURNAL: *Genesis: The Journal of Genetics and Development* 27 (1):p22-31  
May, 2000  
MEDIUM: print  
ISSN: 1526-984X  
DOCUMENT TYPE: Article  
RECORD TYPE: Abstract  
LANGUAGE: English  
SUMMARY LANGUAGE: English

...ABSTRACT: proteins (SH3-domain binding protein, G-protein-coupled  
receptor, Ser-Thr protein kinase), cell adhesion molecules (syndecan-4,  
integrin-beta1), and also gene33, H19, SM20, \*IGFBP5\*, MAMA receptor,  
lectin, keratin, beta-tubulin, calreticulin, GRP78, ERp72, MnSOD,  
thioredoxin, and others. Some have previously been associated with kidney  
development and serve as good controls for expected changes, while most  
have not been linked with kidney epithelial cell differentiation. Using  
thin sections of embryonic kidney and labeled \*antisense\* RNA probes, we  
applied RNA hybridization to confirm the results of DD and related the  
expression of these genes to specific cell lineages of the...

3/3,K/2 (Item 2 from file: 5)  
DIALOG(R)File 5:Biosis Previews(R)  
(c) 2002 BIOSIS. All rts. reserv.

11428292 BIOSIS NO.: 11428292  
Up-regulation of insulin-like growth factor binding protein-5 is  
independent of muscle cell differentiation, sensitive to rapamycin, but  
insensitive to wortmannin and LY294002.

AUTHOR: Rousse Sophie; Montarras Didier; Finsat Christian; Dubois Catherine  
(a)  
AUTHOR ADDRESS: Institut National de la Santé et de la Recherche Médicale, U114, Hôp. Saint  
Louis, 107 Boulevard de l'Hôpital, 75653 Paris Cedex 13 France  
JOURNAL: *Journal of Cell Biology* 150(4):611-618 April, 2000  
MEDIUM: print  
DOCUMENT TYPE: Article  
RECORD TYPE: Article  
LANGUAGE: English

...ABSTRACT: insulin-like growth factor binding protein-5 (IGFBP5) is secreted by the  
cells. The release of myofibroblasts is likely to be associated with a variety  
of factors. IGF-1R transduction RNA was used to show that specific IGFBP5  
expression correlates with the release of the muscle cell lineages of IGF-1R.  
The release of myofibroblasts is associated with the expression of IGFBP5.

SEARCH & FILE NUMBER: SPANISH  
SEARCHED, INDEXED, ABSTRACTED, SERIALIZED, FILED, COMPUTERIZED.

134125934 CA: 134(10)125934z PATENT

IGFBP-5 antisense oligodeoxynucleotide therapy for hormone-regulated tumors

INVENTOR: ALEXANDER, S. G.; RODRIGUEZ, M. M.

LOCATION: U.S.A.

ASSIGNEE: The University of British Columbia; Miyake, Hisamaki

PATENT: PCT International; WO 200101471 A1 DATE: 20.02.2001

APPLICATION: WO 200003495 PCT/CA00/003495 (1999.07.13)

PAGES: 45 pp. PCT/EN: PCT/CA00/003495 LANGUAGE: English CLASS: A61K-048/00A

DESIGNATED COUNTRIES: AE; AG; AL; AM; AT; AU; AZ; BA; BB; BG; BR; BY; BZ; CA; CH; CN; CR; CU; CZ; DE; DK; DM; DO; EE; ES; FI; GB; GR; GE; GH; GM; HR; HU; ID; IL; IN; IS; JP; KE; KG; KP; KR; KZ; LS; LK; LS; LT; LU; LV; MA; MD; MG; MK; MN; ME; MM; MU; NE; NL; PL; PT; RO; RU; SD; SE; SG; SI; SK; SL; TM; TR; TT; TW; UA; UG; US; VE; VN; YU; ZA; ZW; AM; AU; BY; KG; KZ; MD; MU; TZ; TM DESIGNATED REGIONS: GH; IM; KR; LS; MW; MC; SU; SL; SZ; TZ; US; ZW; AT; BE; CH; CY; DE; DK; FI; FR; GR; IE; IT; LU; MC; NL; PT; SE; BE; BY; CH; CG; CI; CM; GA; GW; ML; MR; NE; SN; TD; TG

... completed examination records  
File 101: EP and other items  
Show titles; Refs 3,4,5,6  
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File 17:EMBASE 1974-2002/Aug WI  
Ref 17: 2002 Elsevier Science B.V.  
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Ref 20: 2002 The HW Wilson Co.  
File 21:Wilson Appl. Sci. & Test Abs 1953-2002/Jun  
Ref 21: 2002 The HW Wilson Co.  
File 135:NewsRx Weekly Reports 1995-2002/Aug WI  
Ref 22: 2002 NewsRx  
File 143:Biol. & Agric. Index 1963-2002/Jun  
Ref 23: 2002 The HW Wilson Co.  
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Ref 24: 2002 INSPEC  
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File 172:EMBASE Alert 2 02/Aug W2  
Ref 25: 2002 Elsevier Science B.V.  
File 266:EBRIP 2001/Jun  
Ref 26: Comp & dist by NTIS, Int'l Copyright All Rights Res  
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Ref 27: 2002 DECHEMA  
File 357:Derwent Biotech Res. 1982-2002/June WI  
Ref 28: 2002 Thomson Derwent & ISI  
File 358:Current BioTech Abs 1983-2001/Oct  
Ref 29: 2001 DECHEMA  
File 369:New Scientist 1994-2002/Jul W2  
Ref 30: 2002 Reed Business Information Ltd.  
File 370:Science 1966-1999/Jul W3  
Ref 31: 1999 AAAS  
File 389:CA SEARCH(R) 1967-2002/UD=13707  
Ref 32: 2002 AMERICAN CHEMICAL SOCIETY  
File 434:SciSearch(R) Cited Ref: Sci 1974-1989/Dec  
Ref 33: 1989 Inst for Sci Info  
File 441:Environ Rpt 1970-1991/May  
Ref 34: 1991 Int'l. J. Environ. Res. & Public Health  
Ref 35: 1991 Int'l. J. Environ. Res. & Public Health  
File 442:ICR Annual 1970-1991/May  
Ref 36: 1991 ICR International  
File 443:ENVI.Rpt. 1970-1992/Jan  
Ref 37: 1992 International Academy of Water  
File 444:Conference Papers Index 1973-1992/Jul  
Ref 38: 1992 Cambridge Sri Abs  
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Ref 39: 1995 Geology Suppl. 1995  
File 446:Geology 1970-1994/Jan 1995  
Ref 40: 1995 Geology Suppl. 1995

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(c) 2001 Mass. Med. Soc.

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Set	Items	Description
S1	536	IGFBP5 OR (IGF (W) BPS)
S2	13	S1 (S) (ANTISENSE OR RIBOZYME?)
S3	3	RD (unique items)
S4	14	S1 AND (ANTISENSE OR RIBOZYME?)
S5	4	RD (unique items)
S6	1907	INSULIN (W) LIKE (W) GROWTH (W) FACTOR (W) BINDING (W) PRO-TEIN (W) 5
S7	11	S6 (S) (ANTISENSE OR RIBOZYME?)
S8	4	RD (unique items)
S9	42	S6 AND (ANTISENSE OR RIBOZYME?)
S10	17	RD (unique items)

8/3,K/1 (Item 1 from file: 5)

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12679811 FINGER NO.: 20000333813

Castration-induced up-regulation of insulin-like growth factor binding protein-5 potentiates insulin-like growth factor-I activity and accelerates progression to androgen independence in prostate cancer models.

AUTHOR: Miyake Hideaki; Pollak Michael; Gleave Martin E(a)

**AUTHOR ADDRESS:** (a)Division of Prelogy, University of British Columbia,  
P-3, 2733 Heather Street, Vancouver, BC, V5Z 3J5\*Canada

JOURNAL: *Cancer Research* 60 (11):p3058-3064 June 1, 2000

MEDIUM: print

SEN: 0008-5472

DOCUMENT TYPE: Article

RECORD TYPE: Abstract

NEWTON TYPE: NO. 6  
LANGUAGE: English

SUMMARY LANGUAGE: English

ABSTRACT: Although \*insulin\*-like\* \*growth\* \*factor\* \*binding\* \*protein\*-5\* (IGFBP-5) has been shown to be implicated in prostate cancer progression, the functional role of IGFBP-5 in progression to androgen-independence remains largely...

...cells were stably transfected with IGFBP-5 gene, and IGFBP-5-overexpressing LNCaP tumors progressed significantly faster to androgen independence after castration compared with controls.

\*Antisense\* mouse IGFBP-5 oligodeoxynucleotides (ODNs) were then designed that reduced IGFBP-5 expression in Shionogi tumor cells *in vitro* in a dose-dependent and sequence-specific manner. Growth of Shionogi tumor cells was inhibited by \*antisense\* IGFBP-5 ODN treatment in a time- and dose-dependent manner, which could be reversed by exogenous IGF-I.

However, \*antisense\* IGFBP-5 ODN treatment had no additive inhibitory effect on Shionogi tumor cell growth when IGF-I activity was neutralized by anti-IGF-I antibody. \*Antisense\* IGFBP-5 ODN treatment resulted in decreased mitogen-activated protein kinase activity and number of cells

decreased nitrogen activated protein kinase activity and number of cells in the S + G2-M phases of the cell cycle that directly correlated with reduced proliferation rates of fibroblasts from mice. Systemic administration of the inhibitor of PKC- $\alpha$ , 17-N-methyl- $\alpha$ -methylglibenclamide, after treatment with 5-fluorouracil significantly reduced the proliferation and the thymidine incorporation in fibroblasts from mice.

and the use of a high intensity and/or a pulsed delivery that the response of prostate cancer to an androgen withdrawal can be enhanced by substances, such as tamoxifen, LHRH-agonist therapy, that target LHR signal transduction.

8/3, K/2 (Item 1 from file: 98)

1966-1967 學年第一學期各科成績

similar like it with different differentiation potential. Differentiation induced differentiation through an insulin like growth factor-dependent mechanism.

James, Raymond  
Stewart, Claire K. H.; Kitzmiller, Peter  
The Journal of Cell Biology / Cell Biol. v. 134 no. 1 May 1996 p. 683-93  
SPECIAL FEATURES: ; 134 (1) ISSN: 0021-9751  
LANGUAGE: English  
COUNTRY OF PUBLICATION: United States

ABSTRACT: The function of 'insulin-like' growth factor 'binding' protein-5 (IGFBP-5) in myogenesis, a process stimulated by IGFs, was studied using C2 myoblasts. Compared with vector-transfected control cells, C2 myoblasts expressing an IGFBP-5...

...extracellular matrix during proliferation and subsequently did not differentiate normally. In contrast, differentiation was premature and more extensive in the presence of an IGFBP-5 'antisense' transcript. The inhibitory effects of high IGFBP-5 expression were overcome by exogenous IGFs. These findings are in agreement with a model in which IGFs...

8/3,K/3 (Item 1 from file: 399)

DIALOG(R)File 399:CA SEARCH(R)  
(c) 2002 AMERICAN CHEMICAL SOCIETY. All rts. reserv.

136321287 CA: 136(21)321287w PATENT

Use of pregnancy-associated plasma protein-A2 (PAPP-A2), a novel insulin-like growth factor-binding protein-5 proteinase, for diagnosis and treatment of fetal abnormalities

INVENTOR(AUTHOR): Osvig, Claus; Overgaard, Michael Toft

LOCATION: Den.

ASSIGNEE: Como Biotech Aps

PATENT: PCT International ; WO 200232953 A2 DATE: 20020425

APPLICATION: WO 2001DK695 (20011019) \*DK 20001571 (20001020) \*US PV241840 (20001020)

PAGES: 113 pp. CODEN: PIXXD2 LANGUAGE: English CLASS: C07K-014/435A  
DESIGNATED COUNTRIES: AE; AG; AL; AM; AT; AU; AU; BA; BB; BG; BE; BY;  
BZ; CA; CH; CN; CO; CR; CU; CZ; DE; DE; DK; DM; DZ; EC; EE; ES;  
FI; FI; GB; GE; GH; GM; HR; HU; ID; IL; IN; IS; JP; KE; KG; KP; KR; KZ;  
LC; LK; LR; LS; LT; LU; LV; MA; MD; MG; MN; MW; MX; MZ; NO; NZ; PH; PL;  
PT; RO; RU; SD; SE; SG; SI; SK; SL; TJ; TM; TR; TT; TZ; UA; UG; US; UU;  
VN; YU; ZA; ZW; AM; AZ; BY; KG; KZ; MD; RU DESIGNATED REGIONAL: GH; GM; KE;  
LS; MW; MZ; SD; SL; SZ; TZ; UG; ZW; AT; BE; CH; CY; DE; DK; ES; FI; FR;  
GB; GR; IE; IT; LU; MC; NL; PT; SE; TR; BF; BJ; CF; CG; CI; CM; GA; GN; GQ;  
GW; ML; MR; NE; SN; TD; TG

8/3,K/4 (Item 2 from file: 399)

DIALOG(R)File 399:CA SEARCH(R)  
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124270541 CA: 124(20)270541s PATENT

Use of antisense nucleic acids/analogs inhibiting growth factor-mediated cell proliferation for treatment of proliferative and/or inflammatory skin disorders

INVENTOR(AUTHOR): Winkler, Michael; Winkler, Michael; Winkler, Michael

LOCATION: Austria

ASSIGNEE: St. Lukas University Hospital, Research Foundation

PATENT: PCT International ; WO 2001636411 DATE: 20011226

APPLICATION: WO 2001043411 (20010411) \*AU 94270541s (20010411)  
PAGES: 115 pp. CODEN: PIXXD2 LANGUAGE: English CLASS: A61K-39/77A;  
A61K-39/77B; A61K-39/77C; A61K-39/77D; A61K-39/77E; A61K-39/77F; A61K-39/77G; A61K-39/77H; A61K-39/77I; A61K-39/77J; A61K-39/77K; A61K-39/77L; A61K-39/77M; A61K-39/77N; A61K-39/77O; A61K-39/77P; A61K-39/77Q; A61K-39/77R; A61K-39/77S; A61K-39/77T; A61K-39/77U; A61K-39/77V; A61K-39/77W; A61K-39/77X; A61K-39/77Y; A61K-39/77Z; A61K-39/77Z1